AMENDMENTS TO THE CLAIMS

 (currently amended) A eompound method to treat conditions benefited by modulation of N-type and/or T-type calcium ion channel activity which method comprises administering to a subject in need of such treatment an effective amount of a compound of the formula:

$$Ar - (X^{1})_{n} - N - X^{2} - C - A$$

$$R^{1}_{m}$$

$$N - X^{2} - C - A$$

$$R^{2}_{1}$$

$$(1)$$

or a pharmaceutically acceptable salt thereof,

wherein Ar is phenyl, a six-membered or five-membered ring which is heteroaromatic or a fused aromatic or heteroaromatic system, each of which may optionally be substituted with one or more non-interfering substituents;

X1 is a linker containing 1-5 members:

n is 0 or 1:

each R¹-R³ is independently a non-interfering non-hydrogen substituent;

each 1 is independently 0-5;

m is 0-4:

X² is a linker comprising a chain of at least 5 members;

A is H, OR, SR, NR₂, or halo wherein R is H or lower alkyl (1-6C); with the proviso that

 (a) Ar is a five- or six-membered heteroaryl substituent or an optionally substituted fused aromatic or heteroaromatic substituent; and/or

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- (b) n is 0: and/or
- (c) m is 1-4; and/or

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(d) X² is alkylene substituted by =O, OR, SR, NR₂ and/or halo; and/or

- (e) X² is a chain of at least 6 members; and/or
- (f) X² contains at least one heteroatom selected from N, S and O; and/or
- (g) A is OR, SR, NR2 or halo, wherein R is H or lower alkyl (1-6C) and/or
- (h) Ar is substituted with at least one t-butyl moiety or at least one substituted alkoxy;
 and/or
 - X1 includes at least one heteroatom selected from O, N and S[[,]];

wherein said condition is selected from the group consisting of stroke, anxiety, over-active bladder, inflammatory bowel disease, head trauma, migraine, chronic pain, neuropathic pain, acute pain, epilepsy, hypertension and cardiac arrhythmia.

(currently amended) The eompound method of claim 1 which wherein the compound of formula (1) is of the formula

$$Ar \longrightarrow het \longrightarrow X^{1'} \longrightarrow N \longrightarrow X^{2} \longrightarrow A \qquad (2)$$

wherein Ar, R^1 - R^3 , l, m, X^2 and A are as defined above, "het" is a heteroatom selected from O, S and N, and wherein X^1 is defined as X^1 but lacking one chain member.

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 (currently amended) The eompound method of claim 1 which wherein the compound of formula (1) is of the formula

$$Ar - (X^1)_n - N - C - X^2 - C - A$$

$$R^1_m - 0$$

$$N - C - X^2 - C - A$$

$$R^3_1 - (3)$$

wherein Ar, X^1 , R^1 - R^3 , l, m, n, A and Ar are defined as above, and X^2 is defined as X^2 but lacking one chain member.

4. (currently amended) The eompound method of claim I which wherein the compound of formula (1) is of the formula

$$Ar \longrightarrow (X^1)_n \longrightarrow N \longrightarrow X^2 \longrightarrow A \qquad (4)$$

wherein R^4 is =0, or a carboxylic acid group or ester or amide thereof and Ar, R^2 - R^3 , X^1 , 1, n, X^2 and A are defined as above.

5. (currently amended) The compound method of claim 1, wherein each non-interfering substituent is independently alkyl (1-10C), alkenyl (2-10C), alkynyl (2-10C), aryl (6-10C), arylalkyl (7-16C) or arylalkenyl (7-16C) each optionally further containing 1-4 heteroatoms (N, O or S) each further optionally substituted, or is independently =O, halo, CF₃, OCF, NO₂, NH₂, OH, or SH wherein S may optionally be oxidized.

(canceled)

 (currently amended) The eompound method of claim 1, wherein each 1 is independently 0 or 1.

(canceled)

- (currently amended) The eompound method of claim 1, wherein X¹ is substituted by
 at the position next to the piperazine ring.
- (currently amended) The eompound method of claim 1, wherein each of R² and R³ is independently alkoxy, halo, or alkyl.
- 11. (currently amended) The eompound method of claim 1, wherein m is 1 and R^1 is =0 or a <u>is</u> carboxylic acid group or ester or amide thereof.
- (currently amended) The eompound method of claim I, wherein Ar is optionally substituted phenyl.
- (currently amended) The eompound method of claim 12, wherein said phenyl is unsubstituted or is substituted by one or more tert-butyl, methoxy, substituted alkoxy, hydroxy and/or halo.
- (currently amended) The eompound method of claim 13, wherein the substituted alkoxy is substituted by an amino group.
- (currently amended) The eompound method of claim 1, wherein Ar is optionally substituted pyrimidyl, pyridyl, benzothiazole, benzimidazole or indole.

 (currently amended) The eompound method of claim 1, which is where the compound of formula (1) is

6,6-Bis-(4-fluoro-phenyl)-1-[4-(2-phenylsulfanyl-ethyl)-piperazin-1-yl]-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(4-fluoro-phenoxy)-ethyl]-piperazine;

1-{4-[2-(Benzo[1,3]dioxol-5-yloxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-

1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(2-phenylsulfanyl-ethyl)-piperazine;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(4-methoxy-phenoxy)-ethyl]-piperazine;

1-{4-[2-(2,4-Difluoro-phenoxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-

one:

6,6-Bis-(4-fluoro-phenyl)-1-[4-(2-phenoxy-ethyl)-piperazin-1-yl]-hexan-1-one;

1-{4-[2-(2,4-Dichloro-phenoxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-

one:

6.6-Bis-(4-fluoro-phenyl)-1-{4-[2-(4-methoxy-phenoxy)-ethyl]-piperazin-1-yl}-hexan-1-

one:

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(2-phenoxy-ethyl)-piperazine;

6,6-Bis-(4-fluoro-phenyl)-1-{4-[2-(3,4,5-trimethoxy-phenoxy)-ethyl]-piperazin-1-yl}-hexan-

1-one:

 $1-\{4-[2-(Benzothiazol-2-ylsulfanyl)-ethyl]-piperazin-1-yl\}-6, 6-bis-(4-fluoro-phenyl)-hexandler (A-fluoro-phenyl)-hexandler (A-fluoro-phenyl$

1-one;

[4-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl}-ethoxy)-2,3,6-trimethyl-phenyll-carbamic acid tert-butyl ester;

4-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-ethoxy)-2,3,6-trimethyl-phenylamine:

 $1\hbox{-}[6,6\hbox{-Bis-}(4\hbox{-}fluoro\hbox{-}phenyl)\hbox{-}hexyl]\hbox{-}4\hbox{-}[2\hbox{-}(2,4\hbox{-}dichloro\hbox{-}phenoxy)\hbox{-}ethyl]\hbox{-}piperazine;$

[2-(4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-ylmethyl}-2,6-di-tert-butyl-phenoxy)-ethyl]-dimethyl-amine;

4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-ylmethyl}-2,6-di-tert-butyl-phenol;

1-[4-(3,5-Di-tert-butyl-4-methoxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one:

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-di-tert-butyl-4-methoxy-benzyl)-piperazine;

 $\label{eq:condition} \{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-(3,5-di-tert-butyl-4-methoxy-phenyl)-methanone:$

1-{4-[3,5-Di-tert-butyl-4-(2-dimethylamino-ethoxy)-benzoyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-Benzo[1,3]dioxol-5-ylmethyl-4-[6,6-bis-(4-fluoro-phenyl)-hexyl]-piperazine;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-di-tert-butyl-benzyl)-piperazine;

 $\label{eq:continuous} $$ \{4-[6,6-Bis-(4-fluoro-phenyl]-hexyl]-piperazin-1-yl}-(3,5-di-tert-butyl-4-hydroxy-phenyl)-methanone:$

1-[4-(3,5-Di-tert-butyl-4-hydroxy-benzyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexanl-one;

1-[4-(3,5-Dibromo-4-hydroxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[4-(3,5-Di-tert-butyl-4-hydroxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one:

1-[4-(3,5-Di-tert-butyl-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(4-tert-butyl-benzyl)-piperazine;

 $1\hbox{-[}6,\!6\hbox{-Bis-(}4\hbox{-fluoro-phenyl)-hexyl]-}4\hbox{-(}9H\hbox{-thioxanthen-}9\hbox{-yl)-piperazine};$

 $2\hbox{-}\{4\hbox{-}[6,\!6\hbox{-Bis-}(4\hbox{-fluoro-phenyl})\hbox{-hexyl}]\hbox{-piperazin-}1\hbox{-yl}\}\hbox{-benzothiazole};$

6,6-Bis-(4-fluoro-phenyl)-1-(4-pyrimidin-2-yl-piperazin-1-yl)-hexan-1-one;

2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyll-piperazin-1-yl}-pyrimidine;

6,6-Bis-(4-fluoro-phenyl)-1-[4-(9H-thioxanthen-9-yl)-piperazin-1-yl]-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic acid ethyl ester;

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6.6-Bis-(4-fluoro-phenyl)-1-{4-[2-(3,4,5-trimethoxy-benzylamino)-ethyl]-piperazin-1-yl}-
hexan-1-one:
       9,9-Bis-(4-fluoro-phenyl)-1-[4-(3,4,5-trimethoxy-benzyl)-piperazin-1-yl]-nonan-1-one;
       (2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyll-piperazin-1-yl}-ethyl)-phenyl-amine;
       1-[9,9-Bis-(4-fluoro-phenyl)-nonyll-4-(3,4,5-trimethoxy-benzyl)-piperazine:
      (4-{4-[Bis-(4-fluoro-phenyl)-methoxyl-butyl}-piperazin-1-yl)-(3,4,5-trimethoxy-phenyl)-
methanone:
       6,6-Bis-(4-fluoro-phenyl)-1-[4-(4-trifluoromethoxy-benzoyl)-piperazin-1-yl]-hexan-2-one;
       1-[4-(4-Bromo-benzovl)-piperazin-1-vl]-6.6-bis-(4-fluoro-phenyl)-hexan-1-one:
       6.6-Bis-(4-fluoro-phenyl)-5-hydroxy-1-[4-(3.4.5-trimethoxy-benzoyl)-piperazin-1-yl]-
hexan-1-one:
       1-{4-[Bis-(4-fluoro-phenyl)-methoxy]-butyl}-4-(3,4,5-trimethoxy-benzyl)-piperazine;
       6,6-Bis-(4-fluoro-phenyl)-6-hydroxy-1-[4-(3,4,5-trimethoxy-benzoyl)-piperazin-1-yl]-
hexan-1-one:
       4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic
acid:
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-1-(3,4,5-trimethoxy-benzyl)-piperazin-2-one;
       1-[6.6-Bis-(4-fluoro-phenyl)-hexanoyl]-4-(3.5-di-tert-butyl-4-methoxy-benzoyl)-piperazin-
2-one
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4,5-trimethoxy-benzoyl)-piperazin-2-one;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-(3,4,5-trimethoxy-benzyl)-piperazin-2-one;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-1-(3,5-di-tert-butyl-4-methoxy-benzyl)-piperazin-2-
one:
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanovl]-1-[2-(4-fluoro-phenoxy)-ethyl]-piperazin-2-one;
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-di-tert-butyl-4-methoxy-benzoyl)-piperazin-2-
one;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-1-(3,5-di-tert-butyl-4-methoxy-benzoyl)-piperazin-
2-one:
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyll-4-(3,5-di-tert-butyl-4-methoxy-benzyl)-piperazin-2-one:
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\label{lem:constraint} \hbox{$1$-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-$4-(3,5-di-tert-butyl-4-hydroxy-benzoyl)-piperazin-2-one;}
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6,6-Bis-(4-fluoro-phenyl)-1-[4-(3,4,5-trimethoxy-benzoyl)-piperazin-1-yl]-hex-5-en-1-one; 1-{4-[2-(3,4-Dimethoxy-phenoxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-

1-one:

acid:

 $1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(3,4-dimethoxy-phenoxy)-ethyl]-piperazine;\\ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic and the statement of the statement$

4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic acid ethyl ester;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4-dimethoxy-benzyl)-piperazine;

1-[4-(3,4-Dimethoxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[2-(Benzo[1,3]dioxol-5-yloxy)-ethyl]-4-[6,6-bis-(4-fluoro-phenyl)-hexyl]-piperazine;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(3,4,5-trimethoxy-phenoxy)-ethyl]-piperazine;

1-(4-Amino-2,3,5-trimethyl-phenoxy)-3-{4-[6,6-bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-propan-2-ol;

1-{4-[3-(4-Amino-2,3,5-trimethyl-phenoxy)-2-hydroxy-propyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-(4-Benzothiazol-2-yl-piperazin-1-yl)-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-bis-trifluoromethyl-benzyl)-piperazine;

1-[4-(3,5-Bis-trifluoromethyl-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-

one;

1-[4-(4-tert-Butyl-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(4-bromo-benzyl)-piperazine;

2-(2-{4-{6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-ethylsulfanyl)-benzothiazole; 6.6-Bis-(4-fluoro-phenyl)-1-{4-(4-hydroxy-3,5-dimethoxy-benzoyl)-piperazin-1-yl}-hexan-

1-one:

4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-ylmethyl}-2,6-dibromo-phenol; 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(4-trifluoromethoxy-benzyl)-piperazine:

 $\label{lem:condition} \ensuremath{(2\mbox{-}\{4\mbox{-}\{6,6\mbox{-}Bis\mbox{-}(4\mbox{-}fluoro\mbox{-}phenyl)\mbox{-}hexyl\}\mbox{-}ethyl)\mbox{-}(3,4,5\mbox{-}trimethoxy\mbox{-}benzyl)\mbox{-}amine:}$

 $1-\{4-[2-(4-Fluoro-phenoxy)-ethyl]-piperazin-1-yl\}-6, 6-bis-(4-fluoro-phenyl)-hexan-1-one;\\$

6,6-Bis-(4-fluoro-phenyl)-1-[4-(2-phenylamino-ethyl)-piperazin-1-yl]-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(2,4-difluoro-phenoxy)-ethyl]-piperazine;

N-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-2-oxo-ethyl)-benzamide;

N-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-2-oxo-ethyl)-4-chloro-

benzamide;

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-methyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-isopropyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-tert-butyl-penzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-fluoro-penzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-benzamide; N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-chloro-$

henzamide

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-methyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-isopropyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-tert-butyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-fluoro-penzamide:$

1-[4-(2-Benzylamino-ethyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

 $1-\{4-[2-(4-Chloro-benzylamino)-ethyl]-piperazin-1-yl\}-6, \\ 6-bis-(4-fluoro-phenyl)-hexan-1-one;$

 $6, 6-B is-(4-fluoro-phenyl)-1-\{4-[2-(4-methyl-benzylamino)-ethyl]-piperazin-1-yl\}-hexan-1-yl-benzylamino\}-final final final$

one:

 $\label{lem:constraint} 6, 6-Bis-(4-fluoro-phenyl)-1-\{4-[2-(4-isopropyl-benzylamino)-ethyl]-piperazin-1-yl\}-hexanl-one:$

 $\label{lem:condition} I-\{4-[2-(4-Fluoro-benzylamino)-ethyl]-piperazin-1-yl\}-6, \\ 6-bis-(4-fluoro-phenyl)-hexan-1-one:$

6,6-Bis-(4-fluoro-phenyl)-1-[4-(1-hydroxy-pyridine-4-carbonyl)-piperazin-1-yl]-hexan-1-one;

4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-ylmethyl}-2,6-dimethoxy-phenol; 9.9-Diphenyl-1-[4-(3.4.5-trimethoxy-benzyl)-piperazin-1-yll-nonan-1-one:

1-[4-(3,5-Di-tert-butyl-4-methoxy-benzoyl)-2-methyl-piperazin-1-yl]-6,6-bis-(4-fluorophenyl)-hexan-1-one;

1-[4-(3,5-Di-tert-butyl-4-methoxy-benzoyl)-3-methyl-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[4-(3,5-Di-tert-butyl-benzoyl)-2-methyl-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexanl-one; or

1-[4-(4-tert-Butyl-benzoyl)-2-methyl-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one, including the pharmaceutically acceptable salts thereof.

17-18, (canceled)

19. (new): A compound of the formula

$$A_{1} - (X^{1})_{n} - N - C - X^{2} - C - A$$

$$B - R^{3}_{1}$$

$$A_{1} - (X^{2})_{n} - N - C - X^{2} - C - A$$

$$A_{2} - (X^{2})_{n} - (X^{2})_{n} - C - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - (X^{2})_{n} - C - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - C - A$$

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$$A_{3} - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - C - A$$

$$A_{3} - (X^{2})_{n} - C - C - A$$

wherein Ar is phenyl, a six-membered or five-membered ring which is heteroaromatic or a fused aromatic or heteroaromatic system, each of which may optionally be substituted with one or more non-interfering substituents;

X1 is a linker containing 1-5 members;

n is 1:

each R1-R3 is independently a non-interfering non-hydrogen substituent;

each I is independently 0-5;

m is 0-4:

X2' is a linker comprising a chain of at least 4 members;

A is H, OR, SR, NR₂, or halo wherein R is H or lower alkyl (1-6C); with the proviso that

- (a) Ar is a five- or six-membered heteroaryl substituent or an optionally substituted fused aromatic or heteroaromatic substituent; and/or
 - (b) n is 0: and/or
 - (c) m is 1-4; and/or
 - (d) X^2 is alkylene substituted by =O, OR, SR, NR₂ and/or halo; and/or
 - (e) X² is a chain of at least 6 members; and/or
 - (f) X² contains at least one heteroatom selected from N, S and O; and/or
 - (g) A is OR, SR, NR₂ or halo, wherein R is H or lower alkyl (1-6C) and/or
 - (h) Ar is substituted with at least one t-butyl moiety or at least one substituted alkoxy;

and/or

X¹ includes at least one heteroatom selected from O. N and S.

20. (new): The compound of claim 19, wherein each non-interfering substituent is independently alkyl (1-10C), alkenyl (2-10C), alkynyl (2-10C), aryl (6-10C), arylalkyl (7-16C) or arylalkenyl (7-16C) each optionally further containing 1-4 heteroatoms (N, O or S) each further optionally substituted, or is independently =O, halo, CF₃, OCF, NO₂, NH₂, OH, or SH wherein S may optionally be oxidized.

- 21. (new): The compound of claim 19, wherein each 1 is independently 0 or 1.
- 22. (new): The compound of claim 19, wherein X^1 is substituted by =O at the position next to the piperazine ring.
- $23. \qquad \text{(new): The compound of claim 19, wherein each of R^2 and R^3 is independently alkoxy, halo, or alkyl.}$
- (new): The compound of claim 19, wherein m is 1 and R¹ is =O or a carboxylic acid group or ester or amide thereof.
 - 25. (new): The compound of claim 19, wherein Ar is optionally substituted phenyl.
- (new): The compound of claim 25, wherein said phenyl is unsubstituted or is substituted by one or more tert-butyl, methoxy, substituted alkoxy, hydroxy and/or halo.
- (new): The compound of claim 26, wherein the substituted alkoxy is substituted by an amino group.
- (new): The compound of claim 19, wherein Ar is optionally substituted pyrimidyl, pyridyl, benzothiazole, benzimidazole or indole.
 - 29. (new): A compound selected from the group consisting of 6,6-Bis-(4-fluoro-phenyl)-1-[4-(2-phenylsulfanyl-ethyl)-piperazin-1-yl]-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(4-fluoro-phenoxy)-ethyl]-piperazine; 1-[4-[2-(Benzo[1,3]dioxol-5-yloxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-

1-one:

1-one:

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1-[6.6-Bis-(4-fluoro-phenyl)-hexyl]-4-(2-phenylsulfanyl-ethyl)-piperazine;
       1-[6.6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(4-methoxy-phenoxy)-ethyl]-piperazine;
       1-{4-[2-(2,4-Difluoro-phenoxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-
one;
       6,6-Bis-(4-fluoro-phenyl)-1-[4-(2-phenoxy-ethyl)-piperazin-1-yl]-hexan-1-one;
       1-{4-[2-(2,4-Dichloro-phenoxy)-ethyll-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-
one:
       6,6-Bis-(4-fluoro-phenyl)-1-{4-[2-(4-methoxy-phenoxy)-ethyl]-piperazin-1-yl}-hexan-1-
one;
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(2-phenoxy-ethyl)-piperazine;
       6.6-Bis-(4-fluoro-phenyl)-1-{4-[2-(3.4.5-trimethoxy-phenoxy)-ethyl]-piperazin-1-yl}-hexan-
1-one:
       1-{4-[2-(Benzothiazol-2-ylsulfanyl)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-
1-one:
       [4-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexanovl]-piperazin-1-yl}-ethoxy)-2,3,6-trimethyl-
phenyll-carbamic acid tert-butyl ester:
       4-(2-{4-[6.6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-ethoxy)-2,3.6-trimethyl-
phenylamine;
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(2,4-dichloro-phenoxy)-ethyl]-piperazine;
       [2-(4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-ylmethyl}-2,6-di-tert-butyl-
phenoxy)-ethyll-dimethyl-amine:
       4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-vlmethyl}-2,6-di-tert-butyl-phenol;
       1-[4-(3.5-Di-tert-butyl-4-methoxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-
hexan-1-one:
       1-[4-(3,5-Di-tert-butyl-4-methoxy-benzyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-
```

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-di-tert-butyl-4-methoxy-benzyl)-piperazine;

{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-(3,5-di-tert-butyl-4-methoxy-phenyl)-methanone:

1-{4-{3,5-Di-tert-butyl-4-(2-dimethylamino-ethoxy)-benzoyl}-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-one:

1-Benzo[1,3]dioxol-5-ylmethyl-4-[6,6-bis-(4-fluoro-phenyl)-hexyl]-piperazine;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-di-tert-butyl-benzyl)-piperazine;

{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-(3,5-di-tert-butyl-4-hydroxy-phenyl)-methanone:

1-[4-(3,5-Di-tert-butyl-4-hydroxy-benzyl)-piperazin-1-yl]-6, 6-bis-(4-fluoro-phenyl)-hexan-1-one:

1-[4-(3,5-Dibromo-4-hydroxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

1-[4-(3,5-Di-tert-butyl-4-hydroxy-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

 $1\hbox{-}[4\hbox{-}(3,5\hbox{-}Di\hbox{-}tert\hbox{-}butyl\hbox{-}benzoyl)\hbox{-}piperazin\hbox{-}1\hbox{-}yl]\hbox{-}6,6\hbox{-}bis\hbox{-}(4\hbox{-}fluoro\hbox{-}phenyl)\hbox{-}hexan\hbox{-}1\hbox{-}one;$

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(4-tert-butyl-benzyl)-piperazine;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(9H-thioxanthen-9-yl)-piperazine;

 $2\hbox{-}\{4\hbox{-}[6,\!6\hbox{-Bis-}(4\hbox{-fluoro-phenyl})\hbox{-hexyl}]\hbox{-piperazin-}1\hbox{-yl}\}\hbox{-benzothiazole};$

6,6-Bis-(4-fluoro-phenyl)-1-(4-pyrimidin-2-yl-piperazin-1-yl)-hexan-1-one;

2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-pyrimidine;

6,6-Bis-(4-fluoro-phenyl)-1-[4-(9H-thioxanthen-9-yl)-piperazin-1-yl]-hexan-1-one;

1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic acid ethyl ester:

6,6-Bis-(4-fluoro-phenyl)-1-{4-[2-(3,4,5-trimethoxy-benzylamino)-ethyl]-piperazin-1-yl}-hexan-1-one:

9,9-Bis-(4-fluoro-phenyl)-1-[4-(3,4,5-trimethoxy-benzyl)-piperazin-1-yl]-nonan-1-one;

(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-ethyl)-phenyl-amine;

 $1\hbox{-}[9,9\hbox{-Bis-}(4\hbox{-fluoro-phenyl})\hbox{-nonyl}]\hbox{-}4\hbox{-}(3,4,5\hbox{-trimethoxy-benzyl})\hbox{-piperazine};$

```
(4-{4-[Bis-(4-fluoro-phenyl)-methoxyl-butyl}-piperazin-1-yl)-(3,4,5-trimethoxy-phenyl)-
methanone:
       6.6-Bis-(4-fluoro-phenyl)-1-[4-(4-trifluoromethoxy-benzoyl)-piperazin-1-yll-hexan-2-one:
       1-[4-(4-Bromo-benzovl)-piperazin-1-vl]-6,6-bis-(4-fluoro-phenvl)-hexan-1-one;
       6.6-Bis-(4-fluoro-phenyl)-5-hydroxy-1-[4-(3.4.5-trimethoxy-benzoyl)-piperazin-1-yl]-
hexan-1-one:
       1-{4-[Bis-(4-fluoro-phenyl)-methoxy]-butyl}-4-(3,4,5-trimethoxy-benzyl)-piperazine;
       6,6-Bis-(4-fluoro-phenyl)-6-hydroxy-1-[4-(3,4,5-trimethoxy-benzoyl)-piperazin-1-yl]-
hexan-1-one
       4-[6.6-Bis-(4-fluoro-phenyl)-hexyll-1-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic
acid;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-1-(3,4,5-trimethoxy-benzyl)-piperazin-2-one;
       1-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-4-(3,5-di-tert-butyl-4-methoxy-benzoyl)-piperazin-
2-one:
       1-[6.6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3.4.5-trimethoxy-benzoyl)-piperazin-2-one;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-(3,4,5-trimethoxy-benzyl)-piperazin-2-one;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-1-(3,5-di-tert-butyl-4-methoxy-benzyl)-piperazin-2-
one:
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanovl]-1-[2-(4-fluoro-phenoxy)-ethyll-piperazin-2-one;
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-di-tert-butyl-4-methoxy-benzoyl)-piperazin-2-
one;
       4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-1-(3,5-di-tert-butyl-4-methoxy-benzoyl)-piperazin-
2-one:
       1-[6,6-Bis-(4-fluoro-phenyl)-hexyll-4-(3,5-di-tert-butyl-4-methoxy-benzyl)-piperazin-2-one;
       1-[6.6-Bis-(4-fluoro-phenyl)-hexanovll-4-(3.5-di-tert-butyl-4-hydroxy-benzoyl)-piperazin-2-
one;
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6,6-Bis-(4-fluoro-phenyl)-1-[4-(3,4,5-trimethoxy-benzoyl)-piperazin-1-yl]-hex-5-en-1-one;

- 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(3,4-dimethoxy-phenoxy)-ethyl]-piperazine;
 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic
 acid:
- 4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-(3,4,5-trimethoxy-benzyl)-piperazine-2-carboxylic acid ethyl ester;
 - 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,4-dimethoxy-benzyl)-piperazine;
 - 1-[4-(3.4-Dimethoxy-benzoyl)-piperazin-1-yll-6.6-bis-(4-fluoro-phenyl)-hexan-1-one;
 - 1-[2-(Benzo[1,3]dioxol-5-vloxy)-ethyl]-4-[6,6-bis-(4-fluoro-phenyl)-hexyl]-piperazine;
 - 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(3,4,5-trimethoxy-phenoxy)-ethyl]-piperazine;
- 1-(4-Amino-2,3,5-trimethyl-phenoxy)-3-{4-[6,6-bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-propan-2-ol;
- 1-{4-[3-(4-Amino-2,3,5-trimethyl-phenoxy)-2-hydroxy-propyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;
 - 1-(4-Benzothiazol-2-yl-piperazin-1-yl)-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;
 - 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(3,5-bis-trifluoromethyl-benzyl)-piperazine;
- 1-[4-(3,5-Bis-trifluoromethyl-benzoyl)-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-

one;

- 1-[4-(4-tert-Butyl-benzoyl)-piperazin-1-vl]-6.6-bis-(4-fluoro-phenyl)-hexan-1-one;
- 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(4-bromo-benzyl)-piperazine;
- 2-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-ethylsulfanyl)-benzothiazole;
- 6.6-Bis-(4-fluoro-phenyl)-1-[4-(4-hydroxy-3.5-dimethoxy-benzoyl)-piperazin-1-yl]-hexan-

1-one:

- 4-{4-[6.6-Bis-(4-fluoro-phenyl)-hexyll-piperazin-1-ylmethyl}-2.6-dibromo-phenol;
- 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-(4-trifluoromethoxy-benzyl)-piperazine;
- $(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-ethyl)-(3,4,5-trimethoxy-benzyl)-(3,4,$

amine;

1-{4-[2-(4-Fluoro-phenoxy)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;

 $\label{eq:condition} $$ -(4-fluoro-phenyl)-1-[4-(2-phenylamino-ethyl)-piperazin-1-yl]-hexan-1-one; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-4-[2-(2,4-difluoro-phenoxy)-ethyl]-piperazine; $$ N-(2-[4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl)-benzamide; $$ 1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-2-oxo-ethyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-1-[6,6-Bis-(4-fluoro-phenyl)-h$

N-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-2-oxo-ethyl)-4-chloro-

benzamide:

one:

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-methyl-benzamide;$

N-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl}-2-oxo-ethyl)-4-isopropylbenzamide;

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-tert-butyl-benzamide;$

 $N-(2-\{4-\{6,6-Bis-(4-fluoro-phenyl)-hexyl\}-piperazin-1-yl\}-2-oxo-ethyl)-4-fluoro-penzamide:$

N-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl}-2-oxo-ethyl)-benzamide; N-(2-{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl}-2-oxo-ethyl)-4-chloro-benzamide:

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-methyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-isopropyl-benzamide;$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-tert-butyl-benzamide:$

 $N-(2-\{4-[6,6-Bis-(4-fluoro-phenyl)-hexanoyl]-piperazin-1-yl\}-2-oxo-ethyl)-4-fluoro-penzamide:$

 $\label{lem:condition} 6.6 \cdot Bis \cdot (4 \cdot fluoro-phenyl) - 1 \cdot \{4 \cdot [2 \cdot (4 - methyl \cdot benzylamino) \cdot ethyl] - piperazin \cdot 1 \cdot yl\} - hexan-lone:$ one:

6,6-Bis-(4-fluoro-phenyl)-1-{4-[2-(4-isopropyl-benzylamino)-ethyl]-piperazin-1-yl}-hexan-1-one:

- 1-{4-[2-(4-tert-Butyl-benzylamino)-ethyl]-piperazin-1-yl}-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;
- $1-\{4-[2-(4-Fluoro-benzylamino)-ethyl]-piperazin-1-yl\}-6, 6-bis-(4-fluoro-phenyl)-hexan-1-one;$
- $\label{eq:carbonyl} 6, 6-Bis-(4-fluoro-phenyl)-1-[4-(1-hydroxy-pyridine-4-carbonyl)-piperazin-1-yl]-hexan-1-one;$
 - 4-{4-[6,6-Bis-(4-fluoro-phenyl)-hexyl]-piperazin-1-ylmethyl}-2,6-dimethoxy-phenol; 9,9-Diphenyl-1-[4-(3,4,5-trimethoxy-benzyl)-piperazin-1-yl]-nonan-1-one;
- 1-[4-(3,5-Di-tert-butyl-4-methoxy-benzoyl)-2-methyl-piperazin-1-yl]-6,6-bis-(4-fluorophenyl)-hexan-1-one;
- 1-[4-(3,5-Di-tert-butyl-4-methoxy-benzoyl)-3-methyl-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one;
- 1-[4-(3,5-Di-tert-butyl-benzoyl)-2-methyl-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexanl-one;
- 1-[4-(4-tert-Butyl-benzoyl)-2-methyl-piperazin-1-yl]-6,6-bis-(4-fluoro-phenyl)-hexan-1-one, and the pharmaceutically acceptable salts thereof.
- (new): A pharmaceutical composition which comprises, in admixture with a
 pharmaceutically acceptable excipient, a unit dosage amount of at least one compound of claim 19.
- (new): A pharmaceutical composition which comprises, in admixture with a
 pharmaceutically acceptable excipient, a unit dosage amount of at least one compound of claim 29.